

In the Claims:

Kindly cancel claim 21 without prejudice to the filing of any appropriate continuation applications.

Please amend the claims as follows:

2. (Amended) The compound of claim 1 wherein the base sequence binds to the target portion of the nucleic acid in a manner to inhibit the expression of angiogenin.

3. (Amended) The compound of claim 2 wherein the oligonucleotide analog comprises a modification selected from the group consisting of a modified internucleotide linkage, a modified purine or pyrimidine moiety, a modified sugar moiety, a modified 5' hydroxyl moiety, a modified 3' hydroxyl moiety and a modified 2' hydroxyl moiety.

4. (Amended) The compound of claim 3 wherein the modified internucleotide linkage comprises a substituent having an improved aqueous or lipid solubility or improved resistance to nuclease digestion as compared to an unmodified compound.

5. (Amended) The compound of claim 4 wherein the modified internucleotide linkage is selected from the group consisting of phosphorothioate, N-alkyl phosphoramidates, cycloalkyl phosphoramidates, alkyl phosphonates, cycloalkyl phosphonates, phosphodiester, phosphotriester, C₁ - C₄ alkyl, cycloalkyl, short chain heteroatomic backbone, short chain heterocyclic backbone, morpholino backbone, polyprotein-nucleic acid backbone, peptide-

4 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31 32 33 34 35 36 37 38 39 40 41 42 43 44 45 46 47 48 49 50 51 52 53 54 55 56 57 58 59 60 61 62 63 64 65 66 67 68 69 70 71 72 73 74 75 76 77 78 79 80 81 82 83 84 85 86 87 88 89 90 91 92 93 94 95 96 97 98 99 100

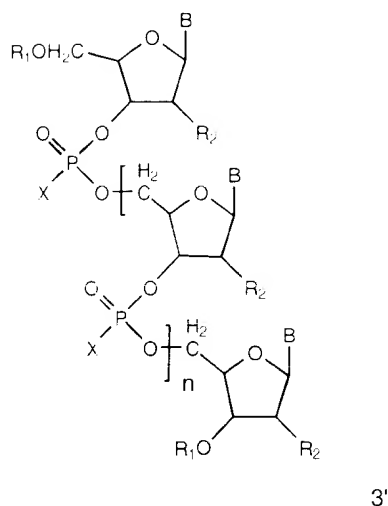
nucleic acid backbone, polyamide, $\text{CH}_2\text{-NH-O-CH}_2$, $\text{CH}_2\text{-N(CH}_3\text{)-O-CH}_2$, $\text{CH}_3\text{-O-N(CH}_3\text{)-CH}_2$, $\text{CH}_2\text{-N(CH}_3\text{)-N(CH}_3\text{)-CH}_2$ and $\text{O-N(CH}_3\text{)-CH}_2\text{-CH}_2$.

8. (Amended) The compound of claim 3 wherein the modified 5' or 3' hydroxyl moiety is selected from the group consisting of C_{1-4} alkoxy, intercalating agent, peptide, enzyme, and ribozyme.

1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31 32 33 34 35 36 37 38 39 40 41 42 43 44 45 46 47 48 49 50 51 52 53 54 55 56 57 58 59 60 61 62 63 64 65 66 67 68 69 70 71 72 73 74 75 76 77 78 79 80 81 82 83 84 85 86 87 88 89 90 91 92 93 94 95 96 97 98 99 100

9. (Amended) The compound of claim 3 wherein the modified 2' hydroxyl moiety is selected from the group consisting of OH, SH, SCH_2 , OCH_3 , F, OCN, OCH_2CH_3 , OCH_2OCH_3 , $\text{OCH}_2\text{O(CH}_2\text{)}_n\text{CH}_3$, $\text{O(CH}_2\text{)}_n\text{NH}_2$, $\text{O(CH}_2\text{)}_n\text{CH}_3$, where n is from 1 to about 10; C_1 to C_{10} lower alkyl, substituted lower alkyl, substituted lower alkaryl substituted lower aralkyl; Cl; Br; CN; CF_3 , OCF_3 , O, S, N-alkyl; O, S, N-alkenyl; SOCH_3 ; SO_2CH_3 ; ONO_2 ; NO_2 ; N_3 ; NH_2 ; heterocycloalkyl, alkaryl; aminoalkylamino; polyalkylamino; substituted silyl; an RNA cleaving group; a cholesteryl group; a conjugate; a reporter group; an intercalator; a group for improving the pharmacokinetic properties of an oligonucleotide as compared to an unmodified compound; and a group for improving the pharmacodynamic properties of an oligonucleotide as compared to an unmodified compound.

5.



X is selected from the group consisting of O, S, and C₁₋₄ alkyl;

R₁ is selected from the group consisting of H, C₁₋₄ alkyl, intercalating agent, peptide, e, and ribozyme;

R₂ is selected from the group consisting of H, OH, SH, SCH₂, OCH₃, F, OCN, OCH₆CH₃, OCH₃OCH₃, OCH₃O(CH₂)_p CH₃, O(CH₂)_pNH₂, O (CH₂)_pCH₃, where p is from 1 to about 10; C₁ to C₁₀ lower alkyl, substituted lower alkyl, substituted lower alkaryl, substituted lower aralkyl; Cl; Br; CN; CF₃; OCF₃; O, S, N-alkyl; O, S, N-alkenyl; SOCH₃; SO₂CH₃; ONO₂; NO₂; N₃; NH₂; heterocycloalkyl, alkaryl; aminoalkylamino; polyalkylamino; substituted silyl; an RNA cleaving group; a cholesteryl group; a conjugate; a reporter group; an intercalator; a group for improving the pharmacokinetic properties of an oligonucleotide as compared to an

unmodified oligonucleotide; and a group for improving the pharmacodynamic properties of an oligonucleotide as compared to an unmodified oligonucleotide; and

n is 5 to 100.

Please add the following new claims:

22. (NEW) The compound of claim 5 wherein the phosphorothioate is selected from the group consisting of alkyl phosphorothioate, cycloalkyl phosphorothioate, and phosphorodithioates.

23. (NEW) The compound of claim 8 wherein the intercalating agent is a substituted acridine.

24. (NEW) The compound of claim 13 wherein the intercalating agent is a substituted acridine.

25. (NEW) The compound of claim 23 wherein the substituted acridine is selected from the group consisting of 2-methoxy-6-chloro-9-pentylaminoacridine, N-(6-chloro-2-methoxyacridinyl)-O-methoxydiisopropylaminophosphinyl-3-aminopropanol, and N-(6-chloro-2-methoxyacridinyl)-O-methoxydiisopropylaminophosphinyl-5-aminopentanol.

26. (NEW) The compound of claim 24 wherein the substituted acridine is selected from the group consisting of 2-methoxy-6-chloro-9-pentylaminoacridine, N-(6-chloro-2-

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methoxyacridinyl)-O-methoxydiisopropylaminophosphinyl-3-aminopropanol, and N-(6 chloro-
2-methoxyacridinyl)-O-methoxydiisopropylaminophosphinyl-5-aminopentanol. --